

**Supplementary information, Fig. S5.**

**(a, b)** Effects of knocking down DR6 (a) or GSDMC (b) on tumor size and weight. DR6 or GSDMC was knocked down in HeLa cells, which were then injected subcutaneously into the posterior flanks of nude mice (n=8). The mice were treated with DM- $\alpha$ KG (500 mg/kg) every other day for 2 weeks. Images of xenograft tumors in the nude mice and tumor weights are shown. PBS was used as a control.

**(c)** The cleavage of GSDMC was determined in tumor samples obtained from (a).

**(d, e)** Comparison of the effects of DR6<sup>WT</sup> and DR6<sup>5CS</sup> on tumor growth (d) and GSDMC cleavage (e). DR6 was knocked down first in HeLa cells, and then DR6<sup>WT</sup> and DR6<sup>5CS</sup> were transfected into cells. Other treatment was the same as (a) and (c).

**(f)** Comparison of GSDMC<sup>WT</sup> and GSDMC<sup>D240A</sup> on tumor growth. GSDMC was knocked down first in HeLa cells, and then GSDMC<sup>WT</sup> and GSDMC<sup>D240A</sup> were transfected into cells. Other treatment was the same as (a).

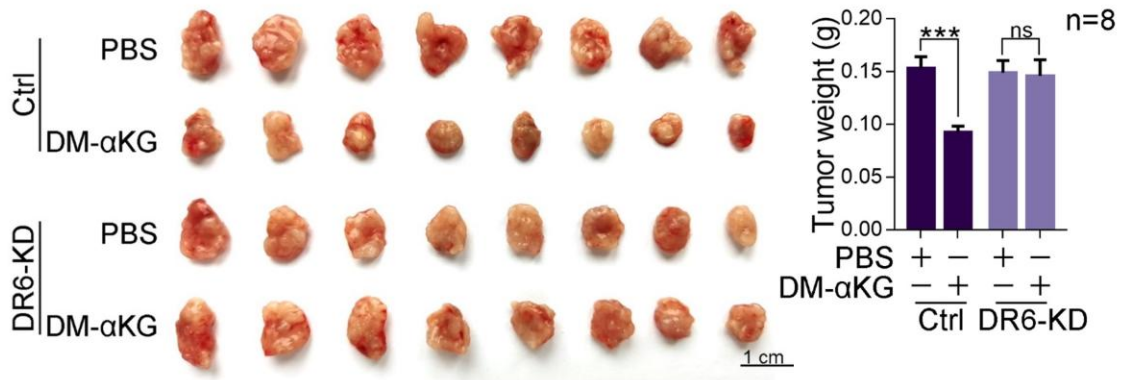
**(g, h, i)** Administration of DM- $\alpha$ KG showed no side effects in the normal C57BL/6J mice. Mice were intraperitoneally injected with DM- $\alpha$ KG (500 mg/kg) or PBS control once per day for 7 days. Mouse body weights (g), images of mouse colon (h) and spleen (i) with colon lengths and spleen weights were indicated.

Tubulin was used to determine the amount of loading proteins. All data are presented as the mean $\pm$ SEM of two or three independent experiments. \*\*  $p < 0.01$ , \*\*\*  $p < 0.001$ , ns: not significant. The data were analyzed using two-tailed Student's t-test in (g, h, i) or

two-way ANOVA followed by the Bonferroni test in (a, b, d, f).

## Supplementary information, Figure S5

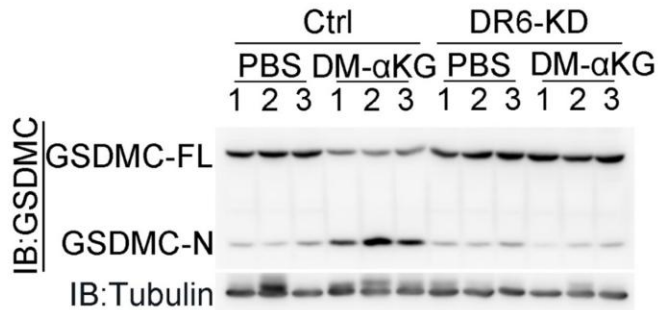
**a**



**b**

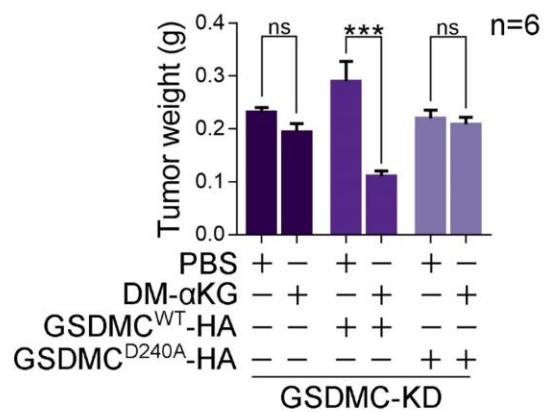
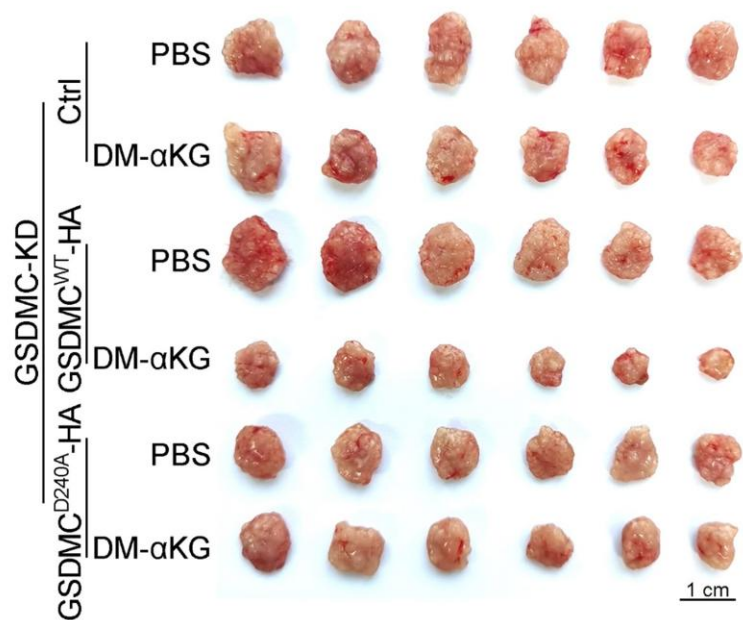


**c**

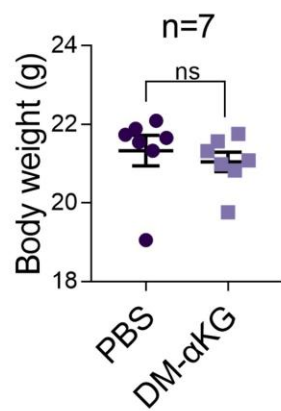




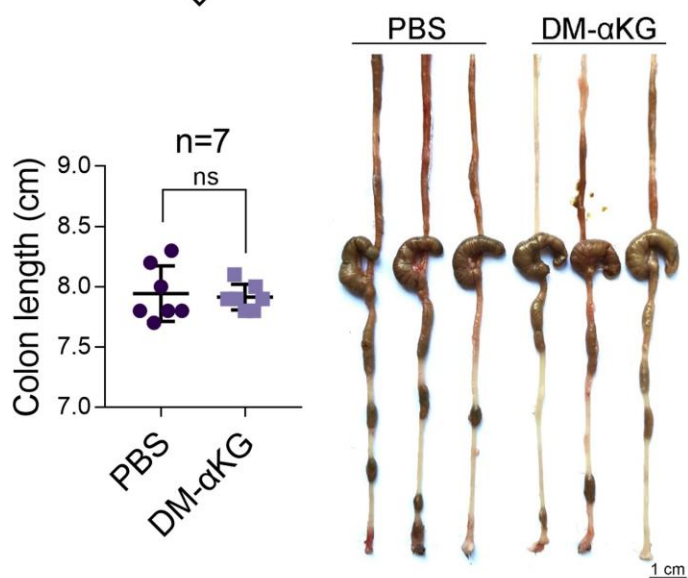
**f**



**g**



**h**



**i**

